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### UTILITY PATENT APPLICATION **TRANSMITTAL**

(Only for new nonprovisional applications under 37 CFR 1.53(b))

Attorney Dock	et No.	05871.0010.CNUS03
First Inventor		Thomas M. Brennan
Title	Metho	od and Apparatus for Performing Numbers of Reactions
Express Mail		EL615212541US

APPLICATION ELE	MENTS		Commissioner for Patents  ADDRESS TO: Box Patent Application									
See MPEP Chapter 600	concerning utility patent app	olication contents.	Washington, DC 20231									
Applicant claims s     See 37 CFR 1.27.     Specification     (preferred arranger     -Descriptive title of     -Cross reference to     -Statement Regard     -Reference to sequence of sequence	al, and a duplicate for fee procesmall entity status.  [Total Pages ment set forth below) the invention a Related Applications ing Fed Sponsored R & D ence listing, a table, gram listing appendix Invention the Invention	ssing) 84 ]	7. ☐ CD-ROM or CD-R in duplicate, large table or Computer Program (Appendix)  8. ☒ Nucleotide and/or Amino Acid Sequence Submission (if applicable, all necessary)  a. ☐ Computer Readable Form (CRF)  b. Specification Sequence Listing on:  i. ☐ CD-ROM or CD-R (2 copies); or  ii. ☐ paper  c. ☒ Statements verifying identity of above copies									
Abstract of the Dis												
5.  Oath or Declaration a.  Newly exect b.  Copy from a (for continuation i.  DELET Signed son amed in 1.63(d)(2 continuation Data states of the continuation of the conti	uted (original or copy) a prior application (37 CFR 1.63 ational divisional with Box 17 co (ION OF INVENTOR(S)) statement attached deleting invinithe prior application, see 37 Cf) and 1.33(b). Sheet. See 37 CFR 1.76  PLICATION, check appropriate der 37 CFR 1.76:  Divisional  Continua	d(d)) impleted) entor(s)  CFR  box, and supply the requition-in-part (CIP) siew Group	9. Assignment papers (cover sheet & document(s))  10. 37 CFR 3.73(b) Statement Power of (when there is an assignee) Attorney  11. English Translation Document (if applicable)  12. Information disclosure Copies of IDS Statement (IDS)/PTO-1449 Citations  13. Preliminary Amendment  14. Return Receipt Postcard (MPEP 503) (Should be specifically itemized)  15. Certified Copy of Priority Document(s) (if foreign priority is claimed)  16. Other:									
For CONTINUATION OR I	DIVISIONAL APPS only: The part of the disclosure of the a	entire disclosure of the	prior application, from which	n an oath or dec	claration is supplied under							
The incorporation can or	nly be relied upon when a por	tion has been inadverter	ntly omitted from the submitt	ed application	parts.							
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NAME (Print/Type	Albert P	Halluin/Wallace Wu	Registration No. 25,227/45,	380								
Signature	M	leth		Date	Signature Date June 23, 2003							

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1. PTO/SB/17 (09-00)
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## FEE TRANSMITTAL For FY 2002

Patent fees are subject to annual revision.

(\$) 551.00

Complete if Known					
Application Number	Not yet Assigned				
Filing Date	June 23, 2003				
First Named Inventor	Thomas M. Brennan				
Examiner Name	Jeffrey Siew				
Group Art Unit	1637				
Attorney Docket No.	05871.0010.CNUS03				

METHOD OF PAYMENT					FEE CALCULATION (continued)							
The Commissioner is hereby authorized to charge indicated fees and credit any overnavments to:					3. ADDITIONAL FEES							
Deposit Acct. No.	posit					Large Fee	Entity Fee	Small Fee	Entity Fee	Fee Description	Fee Paid	
Deposit Hanney Simon Amadel 9 M/hita 11 D					105	130	205	65	Surcharge - late filing fee or oath			
Account Name Howrey Simon Arnold & White, LLP							127	50	227	25	Surcharge – late provisional filing fee or cover sheet	
Charge Any Additional Fee Required							139	130	139	130	Non-English specification	
Under 37 CFR 1.16 and 1.17							147	2,520	147	2,520	For filing a request for ex parte reexamination	
Applicant claims small entity status. See 37 CFR 1.27					112	920*	112	920*	Requesting publication of SIR prior to Examiner action			
2. 🛛 Payment Enclosed:					113	1,840*	113	1,840*	Requesting publication of SIR after Examiner action	<del></del>		
⊠ Ch	☐ Check ☐ Credit card ☐ Money ☐ Other					115	110	215	55	Extension for reply within first month		
	Order  EEE CALCIII ATION					116	400	216	200	Extension for reply within second month		
FEE CALCULATION  1. BASIC FILING FEE					117	920	217	460	Extension for reply within third month			
1 BASIC	Entity	NG FEE Small	Entity				118	1,440	218	720	Extension for reply within fourth month	
Fee Code	Fee	Fee Code	Fee	Fee	Description	Fan Dairl	128	1,960	228	980	Extension for reply within fifth month	
101	<b>(\$)</b> 750	201	( <b>\$</b> ) 375	Litility	filing fee	375.00	119	320	219	160	Notice of Appeal	
106	330	206	165	-	ning ree	3/5.00	120	320	220	160	Filing a brief in support of an appeal	
107	510	207	255	-	iling fee	<b></b>	121	280	221	140	Request for oral hearing	
108	740	208	370		ie filing fee		138	1,510	138	1,510	Petition to institute a public use proceeding	
114	160	214	80		ional filing fee		140	110	240	55	Petition to revive – unavoidable	
	100		00			·	141	1,280	241	640	Petition to revive – unintentional	
					142	1,280	242	640	Utility issue fee (or reissue)			
	SUBTOTAL (1) (\$) 375.00				143	460	243	230	Design issue fee			
2. EXTR	A CLA	IM FE					144	620	244	310	Plant issue fee	
			Extra Claims		Fee from below	Fee Paid	122	130	122	130	Petitions to the Commissioner	
Total Claims	24	-20** =	4	T x	9	] = 36	123	50	123	50	Petitions related to provisional applications	
Independent	$\vdash$		<del></del>	╡		-	126	180	126	180	Submission of Information Disclosure Stmt	
Claims Multiple Depa	1 andent	- 3**=	0	×	140	] = 0	581	40	581	40	Recording each patent assignment per property (times number of properties)	
					140	= 140	146	740	246	370	Filing a submission after final rejection (37 CFR § 1.129(a)	
Large Fee Code	Entity Fee	Small Fee Code	Entity Fee		Fee Descr	iption	149	740	249	370	For each additional invention to be examined (37 CFR § 1.129(b)	
103	(\$) 18	203	(\$) 9	Claims	in excess of 20		179	740	279	370	Request for Continued Examination (RCE)	
102	84	202	42		ndent claims in e	excess of 3	169	900	169	900	Request for expedited examination	
104	280	204	140	-	e dependent clai						of a design application	
109 80 209 40 ** Reissue Independent claims					e (specify)							
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110 18 210 9 ** Reissue claims in excess of 20 and over original patent					]							
	SUBTOTAL (2) (\$)176.00											
**or number previously paid, if greater; For Relssues, see above							1					

SUBMITTED BY	f = f			Complete (if	annlicable)
Name (Print/Type)	Albert P. Hallujn, Wallace Wu	Registration No. (Attorney/Agent)	25,227, 45,380	Telephone	650-463-8109
Signature	We Ma			Date	June 23, 2003

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# METHOD AND APPARATUS FOR PERFORMING LARGE NUMBERS OF REACTIONS USING ARRAY ASSEMBLY

This application claims priority to U.S. Provisional Application Serial

No. 60/158,315, filed October 8, 1999 and to U.S. Application Serial No. 09/686,597, filed October 10, 2000.

#### FIELD OF THE INVENTION

The present invention relates to a method and apparatus for performing a large number of reactions using array assembly. In particular, the present invention features a method and apparatus for performing a large number of chemical and biological reactions by bringing two arrays into close apposition and allowing reactants on the surfaces of two arrays to come into contact. The present invention is exemplified by performing a large number of polynucleotide amplification reactions using array assembly. In addition, the present invention features a method and apparatus for coupling the amplification of polynucleotides and the detection of sequence variations, expression levels, and functions thereof.

### BACKGROUND OF THE INVENTION

Intense efforts are under way to map and sequence the human genome and the genomes of many other species. In June 2000, the Human Genome Project and Celera Genomics announced that a rough draft of the human genome had been completed. This information, however, represents only a reference sequence of the 3-billion-base human genome. The remaining task lies in the determination of sequence variations (e.g., mutations, polymorphisms, haplotypes) and sequence functions, which are important for the study, diagnosis, and treatment of human genetic diseases.

In addition to the human genome, the mouse genome is being sequenced. Genbank provides about 1.2% of the 3-billion-base mouse genome and a rough draft of the mouse genome is expected to be available by 2003 and a finished genome by 2005. The Drosophila Genome Project has also been completed recently. Thus far, genomes of more than 30 organisms have been sequenced.

Traditional nucleic acid sequencing methods include the chemical cleavage method (or the Maxam-Gilbert method) and the chain termination method (or the Sanger method) (Sambrook *et al.*, *Molecular Cloning: A Laboratory Manual*, 2<sup>nd</sup> Ed., Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989). The basic strategy for the chemical cleavage method is to specifically cleave the end-labeled DNA at only one